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(FILE 'HOME' ENTERED AT 15:26:06 ON 08 JUN 2005)

FILE 'HCAPLUS' ENTERED AT 15:26:12 ON 08 JUN 2005

L1 1 US20040127396/PN OR US2000-213995#/AP, PRN

L2 1 (US20040127396 OR CA2312109)/PN OR US2000-213995#/AP, PRN

FILE 'REGISTRY' ENTERED AT 15:28:06 ON 08 JUN 2005

FILE 'HCAPLUS' ENTERED AT 15:28:09 ON 08 JUN 2005

L3 TRA L2 1- RN : 10 TERMS

FILE 'REGISTRY' ENTERED AT 15:28:10 ON 08 JUN 2005

L4 10 SEA L3

FILE 'WPIX' ENTERED AT 15:28:16 ON 08 JUN 2005

L5 1 (US20040127396 OR CA2312109)/PN OR US2000-213995#/AP, PRN

=> b hcap

FILE 'HCAPLUS' ENTERED AT 15:28:41 ON 08 JUN 2005

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FILE COVERS 1907 - 8 Jun 2005 VOL 142 ISS 24

FILE LAST UPDATED: 7 Jun 2005 (20050607/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d all l2 tot/

L2 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2004:533954 HCAPLUS

DN 141:82318

ED Entered STN: 02 Jul 2004

TI Use of furin and furin-like protease inhibitors in the treatment of inflammatory or matrix remodelling diseases

IN Dubois, Claire

PA Can.

SO U.S. Pat. Appl. Publ., 22 pp.

CODEN: USXXCO

DT Patent

LA English

IC ICM A61K038-17

INCL 514002000

CC 1-7 (Pharmacology)

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2004127396	A1	20040701	US 2001-885914	20010622 <--
	CA 2312109	AA	20011223	CA 2000-2312109	20000623 <--

PRAI CA 2000-2312109 A 20000623
 US 2000-213995P P 20000626 <--

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
US 2004127396	ICM	A61K038-17
	INCL	514002000
US 2004127396	NCL	514/002.000
	ECLA	A61K038/57 <--
CA 2312109	ECLA	A61K038/57 <--

AB The present invention provides methods, uses and compns. of an α 1-antitrypsin variant called PDX or a construct, variant, analog, peptide, peptidomimetic, salt, complex or derivative thereof for the treatment of inflammatory or erosive diseases such as rheumatoid arthritis. PDX inhibited collagen-induced arthritis in female Lewis rats.

ST furin protease inhibitor treatment inflammation; matrix remodelling disease treatment furin protease inhibitor; PDX treatment rheumatoid arthritis

IT ~~Peptidomimetics~~
~~(PDX-related)~~ furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT ~~Peptides~~, biological studies
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
~~(PDX-related)~~ furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT Cell proliferation
 (blocking of proprotein convertase-mediated; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT Drug delivery systems
 (carriers, intracellular; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT Disease, animal
 (erosive, treatment of; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT Gene
 RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (for PDX; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT Adenoviral vectors

~~Anti-inflammatory agents~~

~~Antiarthritics~~

~~Antirheumatic agents~~

~~Drug delivery systems~~

~~Gene therapy~~

Human

Mammalia

Transformation, genetic

(furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT Drug delivery systems
 (prodrugs; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT ~~Platelet-derived growth factors~~
 RL: BSU (Biological study, unclassified); BIOL (Biological study)
 (proprotein convertase-mediated endoproteolytic activation of mature, blocking of; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT Extracellular matrix
 (remodelling diseases, treatment of; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)

IT Synovial membrane
 (synoviocyte, recombinant PDX production in rat; furin and furin-like

- protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT ~~inflammation~~
~~Rheumatoid arthritis~~
(treatment of; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT ~~transforming growth factors~~
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(β -), proprotein convertase-mediated endoproteolytic activation of, blocking of; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT ~~Transforming growth factors~~
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(β 1-, PDX inhibition of furin-mediated processing of human; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT 146480-35-5, Gelatinase A
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(PDX inhibition of furin-mediated processing of; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT 9041-92-3DP, α 1-Antitrypsin, PDX mutant
RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(furin inhibitor; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT 9041-92-3D, PDX-mutant, analogs, salts, complexes, derivs.
RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(furin inhibitor; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT 99676-46-7, Proprotein convertase, 141760-45-4, Furin
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(inhibitors; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT 141769-16-3, TACE
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(proprotein convertase-mediated endoproteolytic activation of, blocking of; furin and furin-like protease inhibitors in treatment of inflammatory or matrix remodelling diseases)
- IT 257637-28-8 257904-58-8 476616-83-8 714399-15-2
RL: PRP (Properties)
(unclaimed sequence; use of furin and furin-like protease inhibitors in the treatment of inflammatory or matrix remodelling diseases)

=> b reg

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STRUCTURE FILE UPDATES: 7 JUN 2005 HIGHEST RN 851848-50-5
DICTIONARY FILE UPDATES: 7 JUN 2005 HIGHEST RN 851848-50-5

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*
* The CA roles and document type information have been removed from *
* the IDE default display format and the ED field has been added, *
* effective March 20, 2005. A new display format, IDERL, is now *
* available and contains the CA role and document type information. *
*

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at:
<http://www.cas.org/ONLINE/DBSS/registryss.html>

~~Seq~~ ide l4 tot

L4 ANSWER 1 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN
RN 714399-15-2 REGISTRY
ED Entered STN: 22 Jul 2004
CN Peptide, (Arg-Xaa-Xaa-Arg) (9CI) (CA INDEX NAME)
OTHER NAMES:
CN 4: PN: US20040127396 PAGE: 1 unclaimed sequence
FS PROTEIN SEQUENCE
MF Unspecified
CI MAN
SR CA
LC STN Files: CA, CAPLUS, USPATFULL

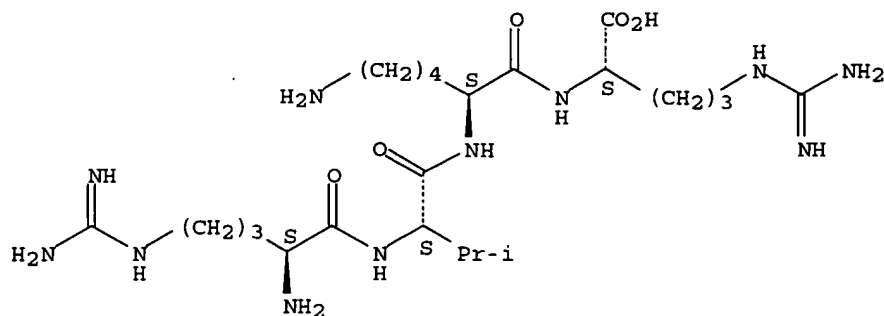
RELATED SEQUENCES AVAILABLE WITH SEQLINK

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
*** USE 'SQD' OR 'SQIDE' FORMATS TO DISPLAY SEQUENCE ***
1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 2 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN
RN 476616-83-8 REGISTRY
ED Entered STN: 18 Dec 2002
CN L-Arginine, L-arginyl-L-valyl-L-lysyl- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN 1: PN: WO2004043385 PAGE: 15 unclaimed sequence
CN 3: PN: US20040127396 SEQID: 3 unclaimed sequence
CN 4: PN: WO02094994 SEQID: 8 unclaimed sequence
CN 52: PN: EP1475435 SEQID: 52 unclaimed sequence
FS PROTEIN SEQUENCE; STEREOSEARCH
MF C23 H47 N11 O5
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.



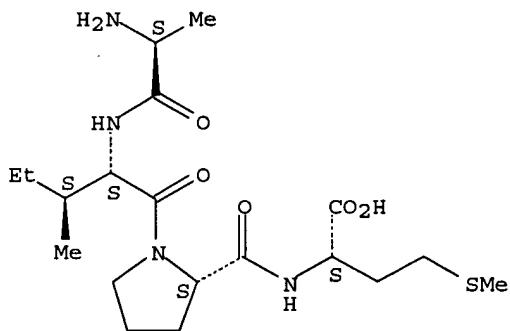
PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

4 REFERENCES IN FILE CA (1907 TO DATE)
4 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 3 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN
RN 257904-58-8 REGISTRY
ED Entered STN: 03 Mar 2000
CN L-Methionine, L-alanyl-L-isoleucyl-L-prolyl- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN 13: PN: US6022855 SEQID: 7 unclaimed sequence
CN 1: PN: US20040127396 SEQID: 1 unclaimed sequence
FS PROTEIN SEQUENCE; STEREOSEARCH
MF C19 H34 N4 O5 S
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

RELATED SEQUENCES AVAILABLE WITH SEQLINK

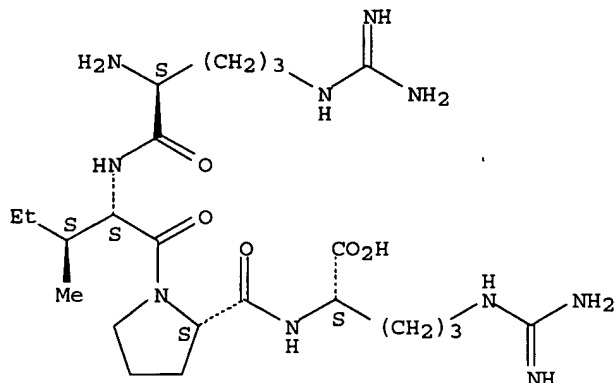
Absolute stereochemistry.



3 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 4 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN
RN 257637-28-8 REGISTRY
ED Entered STN: 01 Mar 2000
CN L-Arginine, L-arginyl-L-isoleucyl-L-prolyl- (9CI) (CA INDEX NAME) PDX
OTHER NAMES:
CN 1: PN: US6022855 SEQID: 10 claimed sequence
CN 2: PN: US20040127396 SEQID: 2 unclaimed sequence
FS PROTEIN SEQUENCE; STEREOSEARCH
MF C23 H44 N10 O5
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

Absolute stereochemistry.



2 REFERENCES IN FILE CA (1907 TO DATE)
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 5 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN
RN 151769-16-3 REGISTRY
ED Entered STN: 16 Dec 1993
CN Proteinase, pro-tumor necrosis factor (9CI) (CA INDEX NAME)
OTHER NAMES:
CN ADAM17 proteinase
CN Metalloprotease TACE
CN Metalloproteinase ADAM17
CN Pro tumor necrosis factor cleavage enzyme
CN Pro-tumor necrosis factor- α -processing enzyme
CN **TACE**
CN TACE proteinase
CN TNF- α convertase
CN TNF- α converting enzyme
CN TNF- α processing protease
CN Tumor necrosis factor α convertase
CN Tumor necrosis factor- α converting enzyme
MF Unspecified
CI MAN
SR CA
LC STN Files: BIOSIS, BIOTECHNO, CA, CAPLUS, CIN, EMBASE, TOXCENTER, USPAT2, USPATFULL

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

490 REFERENCES IN FILE CA (1907 TO DATE)
5 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
495 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 6 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN
RN 147172-61-0 REGISTRY
ED Entered STN: 23 Apr 1993
CN Aggrecanase (9CI) (CA INDEX NAME)
OTHER NAMES:
CN ADAMTS-2
CN ADAMTS-4
CN Aggrecan-degrading metalloproteinase
CN Aggrecan-degrading metalloproteinase ADAMTS4
CN Aggrecan-degrading metalloproteinase MDTs6
CN Aggrecanase 1
CN Metalloproteinase ADAMTS-2

Search done by Noble Jarrell

CN Metalloproteinase ADAMTS-4
CN Proteinase ADAMTS-2
CN Proteinase ADAMTS-4
ENTE A cartilage proteinase
MF Unspecified
CI MAN
SR CA
LC STN Files: AGRICOLA, BIOSIS, BIOTECHNO, CA, CAPLUS, CEN, CIN, EMBASE,
PROMT, TOXCENTER, USPAT2, USPATFULL

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

283 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
288 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 7 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN
RN 146480-35-5 REGISTRY
ED Entered STN: 17 Mar 1993
CN Gelatinase A (9CI) (CA INDEX NAME)
OTHER NAMES:
CN 72 kDa Gelatinase
CN 72 kDa Gelatinase type A
CN 72,000-Mol.-wt. gelatinase
CN 72,000-Mol.-wt. type IV collagenase
CN Collagenase IV
CN Collagenase type IV
CN E.C. 3.4.24.24
CN Matrix metalloprotease 2
CN Matrix metalloproteinase 2
CN MMP 2
CN Type IV collagen metalloproteinase
CN Type IV collagenase
CN Type IV collagenase/gelatinase
MF Unspecified
CI MAN
SR CA
LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
CA, CAPLUS, CEN, CHEMCATS, CIN, EMBASE, PROMT, TOXCENTER, USPAT2,
USPATFULL

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

5383 REFERENCES IN FILE CA (1907 TO DATE)
14 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
5407 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 8 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN
RN 141760-45-4 REGISTRY
ED Entered STN: 12 Jun 1992
CN Furin (enzyme) (9CI) (CA INDEX NAME)
OTHER NAMES:
CN Furin
CN PACE
CN PACE-furin protease
CN Paired basic amino acid cleaving enzyme
CN Paired basic amino acid converting enzyme
CN Saccharomyces cerevisiae gene QDS1 proteinase
CN Serine proteinase PACE
DR 144131-39-5
MF Unspecified
CI MAN
SR CA
LC STN Files: ADISNEWS, AGRICOLA, BIOBUSINESS, BIOSIS, CA, CAPLUS, CEN,
CIN, IPA, PIRA, PROMT, TOXCENTER, USPAT2, USPATFULL

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

704 REFERENCES IN FILE CA (1907 TO DATE)

9 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

705 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 9 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN

RN 99676-46-7 REGISTRY

ED Entered STN: 12 Jan 1986

CN Kex1a (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Adrenorphin-Gly-generating enzyme

CN Conversion endoprotease PC1

CN Convertase PC1

CN Dibasic endopeptidase

CN Dibasic processing protease

CN Dynorphin A-17 processing enzyme

CN E.C. 3.4.21.61

CN Endopeptidase Krp1

CN Endoprotease PC1/3

CN Endoprotease PC3

CN Endoproteinase Kex2p

CN Gene KEX2 dibasic proteinase

CN Kex 2p proteinase

CN Kex2 endopeptidase

CN Kex2 endoprotease

CN Kex2 endoproteinase

CN Kex2 protease

CN Kex2 proteinase

CN Kex2-like endoproteinase

CN Kex2-like precursor protein processing endoprotease

CN Krp1 endopeptidase

CN Neuroendocrine convertase 1

CN PC1 proteinase

CN PC3 precursor-processing proteinase

CN Proconvertase 1

CN Prohormone convertase

CN Prohormone convertase 1

CN Prohormone convertase 3

CN Prohormone convertase I

CN Prohormone convertase PC1

CN Prohormone convertase PC3

CN Prohormone-processing endoprotease

CN Prohormone-processing KEX2 proteinase

CN Prohormone-processing proteinase

CN Proprotein convertase

CN Proprotein convertase Kex2p

CN Proprotein convertase PC1

CN Protease KEX2

CN Proteinase Kex2p

CN Proteinase PC1

CN Proteinase, prohormone-processing

MF Unspecified

CI MAN

SR CA

LC STN Files: AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, CA, CAPLUS, CIN,
EMBASE, PROMT, TOXCENTER, USPAT2, USPATFULL

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

1071 REFERENCES IN FILE CA (1907 TO DATE)

13 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

1074 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L4 ANSWER 10 OF 10 REGISTRY COPYRIGHT 2005 ACS on STN

RN 9041-92-3 REGISTRY

ED Entered STN: 16 Nov 1984

CN Trypsin inhibitor, α 1- (9CI) (CA INDEX NAME) *Mutant PDX*
OTHER NAMES:
CN α 1-Protease inhibitor
CN α 1-Antiprotease
CN α 1-Antiproteinase
CN α 1-Antitrypsin
CN α 1-Antitrypsin Pittsburgh mutant
CN α 1-Antitrypsin Portland
CN α 1-AT
CN α 1-Protease inhibitor
CN α 1-Proteinase inhibitor
CN α 1-Trypsin inhibitor
CN Antitrypsin Pittsburgh
CN Prolastin
CN Respitin
CN SERPINA1
DR 9082-50-2, 124542-00-3
MF Unspecified
CI COM, MAN
LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS,
BIOSIS, BIOTECHNO, CA, CAPLUS, CBNB, CEN, CHEMCATS, CHEMLIST, CIN,
CSCHEM, DDFU, DIOGENES, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB,
IMSCOSEARCH, IMSRESEARCH, IPA, MRCK*, NIOSHTIC, PHAR, PROMT, RTECS*,
TOXCENTER, USPAT2, USPATFULL
(*File contains numerically searchable property data)
Other Sources: EINECS**
(*Enter CHEMLIST File for up-to-date regulatory information)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

5514 REFERENCES IN FILE CA (1907 TO DATE)
311 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
5525 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> b wpix

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FILE LAST UPDATED: 3 JUN 2005 <20050603/UP>
MOST RECENT DERWENT UPDATE: 200535 <200535/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

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FOR DETAILS. <<<

=> d all 15 tot

L5 ANSWER 1 OF 1 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
AN 2002-395072 [43] WPIX
DNC C2002-111283
TI Treating inflammatory or erosive disease e.g. rheumatoid arthritis
comprises administering alpha-one antitrypsin inhibitor variant.
DC B04
IN DUBOIS, C
PA (UYSH) UNIV SHERBROOKE; (DUBO-I) DUBOIS C
CYC 2
PI CA 2312109 A1 20011223 (200243)* EN 50 A61K038-43 <--
US 2004127396 A1 20040701 (200444) A61K038-17 <--
ADT CA 2312109 A1 CA 2000-2312109 20000623; US 2004127396 A1 Provisional
US 2000-213995P 20000626, US 2001-885914 20010622
PRAI CA 2000-2312109 20000623
IC ICM A61K038-17; A61K038-43
ICS A61K038-16; A61K048-00; A61P019-00; A61P029-00
AB CA 2312109 A UPAB: 20020709
NOVELTY - Treating inflammatory or erosive disease involving furin or
furin-like protease activity comprises administering a compound selected
from PDX (serpin alpha 1-antitrypsin inhibitor variant) or construct,
variant, analog, PDX-related peptide, PDX-related peptidomimetic, their
salts, complexes or derivatives.
ACTIVITY - Antiinflammatory; Antirheumatic; Antiarthritic.
MECHANISM OF ACTION - Furin and furin-like protease inhibitor.
Furin-mediated endoproteolytic activation of TGF (transforming growth
factor), PDGF (platelet derived growth factor), TACE (tumor necrosis
factor (TNF- alpha) converting enzyme), and aggrecanase-1 blocker.
USE - Used for treating inflammatory or erosive disease, e.g.
rheumatoid arthritis (claimed) and matrix remodelling conditions in a
mammal.
ADVANTAGE - PDX and its related compounds mimic the minimum consensus
sequence (R-X-X-R) required for furin recognition and are potent furin
inhibitors in vitro and in cells.
Dwg.0/9
FS CPI
FA AB; DCN
MC CPI: B04-C01; B14-C03; B14-C06; B14-C09

=> b home

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